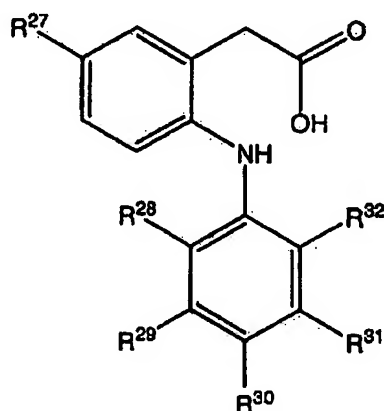


What is claimed is:

1. A combination comprising a cyclooxygenase-2 selective inhibitor in a first amount and a topoisomerase II inhibitor in a second amount, wherein said first amount together with said second amount comprises a therapeutically effective amount for treatment of neoplasia or a neoplasia related disorder, and wherein the cyclooxygenase-2 selective inhibitor is a compound of formula



or an isomer, pharmaceutically acceptable salt, ester, or prodrug thereof, wherein:

$R^{27}$  is methyl, ethyl or propyl;

$R^{28}$  is chloro or fluoro;

$R^{29}$  is hydrogen, fluoro or methyl;

$R^{30}$  is hydrogen, fluoro, chloro, methyl, ethyl, methoxy, ethoxy or hydroxy;

$R^{31}$  is hydrogen, fluoro or methyl; and

$R^{32}$  is chloro, fluoro, trifluoromethyl, methyl or ethyl;

provided that  $R^{28}$ ,  $R^{29}$ ,  $R^{31}$  and  $R^{32}$  are not all fluoro when  $R^{27}$  is ethyl and  $R^{30}$  is hydrogen.

2. The combination of Claim 1 wherein the COX-2 selective inhibitor is lumiracoxib,

3. The combination of Claim 1 or Claim 2 wherein the topoisomerase II inhibitor is selected from the group consisting of aclarubicin, amonafide, amrubicin, amsacrine, annamycin, 6,9-bis[(2-aminoethyl)amino]-benz[g]isoquinoline-5,10-dione, 1,11-dichloro6-[2-(diethylamino)ethyl]-12,13-dihydro-12-(4-O-methyl- $\beta$ -D-glucopyranosyl)-5H-indolo[2,3-a]pyrrolo[3,4-c]carbazole-5,7(6H)-dione, crisnatol, daunorubicin, doxorubicin, epirubicin, etoposide, galarubicin, idarubicin, iododoxorubicin, 10-[[6-deoxy-2-O-(6-deoxy-3-O-methyl- $\alpha$ -D-galactopyranosyl)-3,4-O-[(S)-phenylmethylene]- $\beta$ -D-galactopyranosyl]oxy]-5,12-dihydro-1-methyl-5,12-dioxobenzo[h][1]benzopyrano[5,4,3-cde][1]benzopyran-6-yl ester-3-ethoxypropanoic acid, 8-ethyl-7,8,9,10-tetrahydro-1,6,7,8,11-pentahydroxy-10-[[2,3,6-trideoxy-3-(4-morpholinyl)- $\alpha$ -L-lyxo-hexopyranosyl]oxy]-5,12-naphthacenedione, (7S,9S)-7-[[4-O-(3-amino-2,3,6-trideoxy- $\alpha$ -L-lyxo-hexopyranosyl)-2,6-dideoxy- $\alpha$ -L-lyxo-hexopyranosyl]oxy]-7,8,9,10-tetrahydro-6,9,11-trihydroxy-9-(hydroxyacetyl)-5,12-naphthacenedione, merbarone, mitoxantrone, nemorubicin, pirarubicin, N-[2-(dimethylamino)ethyl]-9-hydroxy-5,6-dimethyl-6H-pyrido[4,3-b]carbazole-1-carboxamide, suboxane, teniposide and valrubicin.

4. The combination of Claim 1 or Claim 2 wherein the topoisomerase II inhibitor is selected from the group consisting of doxorubicin, epirubicin and idarubicin.

5. The combination of any of Claims 1 to 4 wherein the cyclooxygenase-2 selective inhibitor and the topoisomerase II inhibitor are administered substantially simultaneously.

6. The combination of any of Claims 1 to 4 wherein the cyclooxygenase-2 selective inhibitor and the

topoisomerase II inhibitor are administered sequentially.

7. Use of a combination of any of Claims 1 to 4 in manufacture of a medicament for treatment of neoplasia or a neoplasia related disorder.

8. The use of Claim 7 wherein the neoplasia or neoplasia related disorder comprises one or more malignant tumor growths selected from the group consisting of acral lentiginous melanoma, actinic keratoses, acute lymphocytic leukemia, acute myeloid leukemia, adenocarcinoma, adenoid cystic carcinoma, adenomas, adenosarcoma, adenosquamous carcinoma, anal canal cancer, anal cancer, anorectal cancer, astrocytic tumors, bartholin gland carcinoma, basal cell carcinoma, biliary cancer, bone cancer, bone marrow cancer, brain cancer, breast cancer, bronchial cancer, bronchial gland carcinomas, carcinoids, carcinoma, carcinosarcoma, cholangiocarcinoma, chondrocarcinoma, chorioid plexus papilloma/carcinoma, chronic lymphocytic leukemia, chronic myeloid leukemia, clear cell carcinoma, colon cancer, colorectal cancer, connective tissue cancer, cystadenoma, digestive system cancer, duodenum cancer, endocrine system cancer, endodermal sinus tumor, endometrial hyperplasia, endometrial stromal sarcoma, endometrioid adenocarcinoma, endothelial cell cancer, ependymal cancer, epithelial cell cancer, esophageal cancer, Ewing's sarcoma, eye and orbit cancer, female genital cancer, focal nodular hyperplasia, gallbladder cancer, gastric antrum cancer, gastric fundus cancer, gastrinoma, germ cell tumors, glioblastoma, glucagonoma, heart cancer, hemangiblastomas, hemangioendothelioma, hemangiomas, hepatic adenoma, hepatic adenomatosis, hepatobiliary cancer, hepatocellular carcinoma, Hodgkin's disease, ileum cancer, insulinoma, intraepithelial neoplasia, interepithelial squamous cell neoplasia, intrahepatic bile duct cancer, invasive squamous cell carcinoma, jejunum cancer, joint cancer, Kaposi's sarcoma, kidney and renal pelvic cancer, large cell

carcinoma, large intestine cancer, larynx cancer, leiomyosarcoma, lentigo maligna melanomas, leukemia, liver cancer, lung cancer, lymphoma, male genital cancer, malignant melanoma, malignant mesothelial tumors, medulloblastoma, medulloepithelioma, melanoma, meningeal cancer, mesothelial cancer, metastatic carcinoma, mouth cancer, mucoepidermoid carcinoma, multiple myeloma, muscle cancer, nasal tract cancer, nervous system cancer, neuroblastoma, neuroepithelial adenocarcinoma, nodular melanoma, non-epithelial skin cancer, non-Hodgkin's lymphoma, oat cell carcinoma, oligodendroglial cancer, oral cavity cancer, osteosarcoma, ovarian cancer, pancreatic cancer, papillary serous adenocarcinoma, penile cancer, pharynx cancer, pituitary tumors, plasmacytoma, prostate cancer, pseudosarcoma, pulmonary blastoma, rectal cancer, renal cell carcinoma, respiratory system cancer, retinoblastoma, rhabdomyosarcoma, sarcoma, serous carcinoma, sinus cancer, skin cancer, small cell carcinoma, small intestine cancer, smooth muscle cancer, soft tissue cancer, somatostatin-secreting tumor, spine cancer, squamous cell carcinoma, stomach cancer, striated muscle cancer, submesothelial cancer, superficial spreading melanoma, T cell leukemia, testicular cancer, thyroid cancer, tongue cancer, undifferentiated carcinoma, ureter cancer, urethra cancer, urinary bladder cancer, urinary system cancer, uterine cervix cancer, uterine corpus cancer, uveal melanoma, vaginal cancer, verrucous carcinoma, VIPoma, vulva cancer, well differentiated carcinoma and Wilm's tumor.

9. The use of Claim 7 wherein the neoplasia or neoplasia related disorder is selected from the group consisting of lung cancer, colorectal cancer, breast cancer, prostate cancer, bladder cancer, ovary cancer, cervical cancer, gastrointestinal cancer and leukemia.